In the Claims

1. (Canceled) A method for *in situ* formation of a superficial cartilage layer over an articular cartilage lesion, said method comprising steps:

a) obtaining a piece of autologous or heterologous cartilage;

b) isolating, culturing and expanding chondrocytes into a neo-cartilage;

c) preparing a neo-cartilage construct;

d) implanting said neo-cartilage construct into said lesion; and

e) depositing a top biocompatible adhesive sealant over the neo-cartilage construct.

2. (Canceled) The method of claim 1 additionally comprising a step of depositing a biocompatible adhesive sealant at a bottom of the lesion wherein said bottom and top sealant may be the same or different.

3. (Canceled) The method of claim 2 wherein said neo-cartilage construct comprises neo-cartilage chondrocytes embedded into a support matrix.

4. (Currently amended) The method of claim 23 wherein said support matrix is a gel, sponge, honeycomb or lattice ~~a two or three-dimensional structure~~ prepared from a compound selected from the group consisting of a thermo-reversible gelation hydrogel, collagenous gel, Type I collagen, Type II collagen, Type IV collagen, gelatin, agarose, ~~a cell-contracted~~ collagen containing proteoglycan, ~~a cell-contracted~~ collagen containing glycosaminoglycan, ~~a cell-contracted~~ collagen containing glycoprotein, fibronectin, laminin, a growth factor, cytokine, elastin, hyaluronin, fibrin, a synthetic polymeric fiber made of polylactic acid, a synthetic polymeric fiber made of polyglycotic

acid, a synthetic polymeric fiber made of polyamino acid, polycaprolactone, polyamino acid, polypeptide gel, a copolymer[[s]] thereof and a combination thereof.

5. (Original) The method of claim 4 wherein said matrix is the thermo-reversible gelation hydrogel (TRGH) wherein said TRGH is in a liquid sol state at temperatures of below about 30°C and wherein said thermoreversible hydrogel polymer is in a solid sol state at temperature above about 30°C and further wherein said thermo-reversible gelation hydrogel is either deposited into the lesion cavity formed below the top sealant or between the bottom and top sealants as the neo-cartilage construct comprising chondrocytes embedded therein or wherein said TRGH is deposited into said cavity as a space holding gel without any neo-cartilage chondrocytes.

6. (Currently amended) The method of claim 5 wherein the neo-cartilage construct comprises cultured differentiated autologous or heterologous chondrocytes or cells which could be differentiated into chondrocytes~~, said chondrocytes or cells incorporated into said support matrix and subjected to an algorithm of the invention wherein said algorithm comprises constant or cyclic hydrostatic pressure, static atmospheric pressure or non-pressure conditions, perfusion flow rate, medium composition, temperature, cell density, oxygen concentration and time to which the chondrocytes are subjected~~.

7. (Currently amended) The method of claim 24[[5]] wherein said ~~top and~~ bottom sealant is selected from the group consisting of gelatin, a copolymer of polyethylene glycol and poly-lactide or poly-glycolide, periodate-oxidized gelatin, 4-armed pentaerythritol thiol and a polyethylene glycol diacrylate, 4-armed tetra-succinimidyl ester or tetra-thiol derivatized PEG, photo-polymerizable polyethylene glycol-co-poly(α-hydroxy acid)

diacrylate macromer, 4-armed polyethylene glycol derivatized with succinimidyl ester and thiol plus methylated collagen hydrogel, derivatized polyethylene glycol (PEG), derivatized polyethylene glycol (PEG) cross-linked with alkylated collagen, tetra-hydrosuccinimidyl or tetra-thiol derivatized PEG, cross-linked PEG with methylated collagen and a combination thereof ~~and wherein the top and bottom sealants are the same or different~~.

8. (Currently amended) The method of claim 7 wherein the bottom sealant is the cross-linked PEG with methylated collagen.

9. (Original) The method of claim 8 wherein the neo-cartilage construct is prepared *in vitro*, *ex vivo* or *in vivo*.

10. (Canceled) The method of claim 9 wherein the neo-cartilage construct is prepared *ex vivo* and is subjected to an algorithm of the invention.

11. (Canceled) The method of claim 10 wherein the algorithm comprises cyclic or constant hydrostatic pressure, static pressure, flow rate, temperature, time, cell density and oxygen and carbon dioxide content.

12. (Currently amended) The method of claim 23 ~~11~~ wherein the hydrostatic pressure is from about zero MPa to about 10 MPa above atmospheric pressure at about 0.01 to about 1 Hz, wherein the time for applying the hydrostatic pressure is from zero to about 24 hours per day for from about one day to about ninety days, wherein said hydrostatic pressure is preceded or followed by a period of zero to about 24 hours per day of a static atmospheric pressure for from about one day to about ninety days, wherein the flow rate is from about 1 μL/min to about 500 μL/min, wherein the cell density is from about 3 to 60 millions and wherein the oxygen concentration is from about 1 to about 20%.

13. (Original) The method of claim 12 wherein the hydrostatic cyclic pressure is from about 0.05 MPa to about 3 MPa at 0.1 to about 0.5 Hz or constant pressure is from about zero to about 3 MPa above atmospheric pressure and wherein such pressure is applied for about 7 to about 28 days.

14. (Original) The method of claim 13 wherein said hydrostatic pressure is preceded or followed by a period of about zero to about 28 days of atmospheric pressure.

15. (Original) The method of claim 14 wherein said perfusion flow rate is from about 5 μL to about 50 μL/minute.

16. (Original) The method of claim 15 wherein said perfusion flow rate is about 5 μL/minute.

17. (Original) The method of claim 16 wherein said perfusion and pressure are applied at from about 2% to about 5% of oxygen concentration.

18. (Canceled) The method of claim 17 wherein said neo-cartilage construct is implanted into said lesion between said two layers of the sealants.

19. (Currently amended) The method of claim 22 ~~18~~ wherein said neo-cartilage implant is overgrown by said superficial cartilage layer and wherein said superficial cartilage layer and a surrounding native synovial membrane are mutually integrated.

20. (Canceled) The method of claim 19 wherein said neo-cartilage and a surrounding native cartilage are mutually integrated.

21. (Currently amended) The method of claim 5 [[~~20~~]] wherein said neo-cartilage construct comprises a thermo-reversible gelation hydrogel and is implanted into the lesion as a liquid sol wherein upon warming the construct to a body temperature, the liquid sol is converted to a solid gel and wherein this process can be reversed by cooling said lesion to a temperature below 30°C permitting removal of said gel as the sol.

22. (New) A method for treatment of an articulate cartilage lesion and for formation of a superficial cartilage layer, said method comprising surgically implanting a neo-cartilage construct into said lesion and covering said neo-cartilage implant with a layer of a top biocompatible adhesive sealant wherein said top sealant is a derivatized polyethylene glycol (PEG) cross-linked with collagen.

23. (New) The method of claim 22, comprising steps:

a) obtaining an autologous or heterologous cartilage and subjecting said cartilage to a process for isolating of chondrocytes;

b) expanding and suspending said isolated chondrocytes in a gel, sol, sol-gel, collagen or collagen-containing solution;

c) seeding said chondrocytes suspension into a support matrix, wherein said support matrix is a three-dimensional structure containing plurality of pores;

d) preparing said neo-cartilage construct for implantation by subjecting said seeded support matrix to conditions promoting activation and propagation of said chondrocytes within said support matrix wherein said conditions comprise a cyclic or constant hydrostatic pressure, static pressure, medium flow rate, temperature, time, cell

density, oxygen or carbon dioxide content, each alone or in combination;

e) implanting said neo-cartilage construct into said cartilage lesion; and

e) depositing the top biocompatible adhesive sealant over the neo-cartilage construct wherein said top sealant is the polyethylene glycol cross-linked with methylated collagen.

24. (New) The method of claim 23 additionally comprising a step of depositing a layer of a bottom biocompatible adhesive sealant into said cartilage lesion before implanting said neo-cartilage construct, wherein said sealant may be the same as, or different from the top sealant.

25. (New) The method of claim 24 wherein said chondrocytes are isolated from extracellular matter by enzymatic digestion of the cartilage.

26. (New) The method of claim 23 wherein said step of depositing the top sealant over the neo-cartilage construct results in formation of the superficial cartilage layer over the treated lesion.

27. (New) The method of claim 26 wherein said superficial cartilage layer is integrated into a synovial membrane.

28. (New) The method of claim 27 wherein such formation and integration occurs in two or three months following the surgery.